

SEQUENCE LISTING

<110> Kotwal, Girish J. Daly, James

<120> Application of a Viral Complement Inhibitory Protein in the Treatment and Diagnosis of Alzheimer's Disease

<130> 17541/033US1

<140> 09/889,624

<141> 2001-11-07

<150> PCT/US00/01115

<151> 2000-01-19

<150> 60/116,328

<151> 1999-01-19

<160> 12

<170> FastSEQ for Windows Version 4.0

<210> 1

<211> 263

<212> PRT

<213> Vaccinia Virus

<400> 1

Met Lys Val Glu Ser Val Thr Phe Leu Thr Leu Leu Gly Ile Gly Cys

1 10 15

Val Leu Ser Cys Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Lys Phe 20 25 30

Lys Asn Ser Val Glu Thr Asp Ala Asn Ala Asn Tyr Asn Ile Gly Asp 35 40 45

Thr Ile Glu Tyr Leu Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly
50 60

Pro Ile Tyr Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln 65 70 75 80

Cys Ile Lys Arg Arg Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly Gln 85 90 95

Leu Asp Ile Gly Gly Val Asp Phe Gly Ser Ser Ile Thr Tyr Ser Cys
100 105 110

Asn Ser Gly Tyr His Leu Ile Gly Glu Ser Lys Ser Tyr Cys Glu Leu 115 120 125

Gly Ser Thr Gly Ser Met Val Trp Asn Pro Glu Ala Pro Ile Cys Glu 130 135 140

Ser Val Lys Cys Gln Ser Pro Pro Ser Ile Ser Asn Gly Arg His Asn

145 150 155 160

Gly Tyr Glu Asp Phe Tyr Thr Asp Gly Ser Val Val Thr Tyr Ser Cys 165 170 175

Asn Ser Gly Tyr Ser Leu Ile Gly Asn Ser Gly Val Leu Cys Ser Gly
180 185 190

Gly Glu Trp Ser Asp Pro Pro Thr Cys Gln Ile Val Lys Cys Pro His
195 200 205

Pro Thr Ile Ser Asn Gly Tyr Leu Ser Ser Gly Phe Lys Arg Ser Tyr

```
220
   210
                       215
Ser Tyr Asn Asp Asn Val Asp Phe Lys Cys Lys Tyr Gly Tyr Lys Leu
                   230
                                      235
Ser Gly Ser Ser Ser Ser Thr Cys Ser Pro Gly Asn Thr Trp Lys Pro
                                  250
               245
Glu Leu Pro Lys Cys Val Arg
           260
<210> 2
<211> 840
<212> DNA
<213> Vaccinia Virus
tttttattat ttgtacgatg tccaggataa catttttacg gataaataaa tatgaaggtg 60
gagagcgtga cgttcctgac attgttggga ataggatgcg ttctatcatg ctgtactatt 120
ccgtcacgac ccattaatat gaaatttaag aatagtgtgg agactgatgc taatgctaat 180
tacaacatag gagacactat agaatatcta tgtctacctg gatacagaaa gcaaaaaatg 240
ggacctatat atgctaaatg tacaggtact ggatggacac tctttaatca atgtattaaa 300
cggagatgcc catcgcctcg agatatcgat aatggccaac ttgatattgg tggagtagac 360
tttggctcta gtataacgta ctcttgtaat agcggatatc atttgatcgg tgaatctaaa 420
tcgtattgtg aattaggatc tactggatct atggtatgga atcccgaggc acctatttgt 480
qaatctqtta aatqccaatc ccctccatct atatccaacg gaagacataa cggatacgag 540
qatttttata ccqatqqqaq cqttqtaact tatagttgca atagtggata ttcgttgatt 600
ggtaactctg gtgtcctgtg ttcaggagga gaatggtccg atccacccac gtgtcagatt 660
gttaaatgtc cacatcctac aatatcaaac ggatacttgt ctagcgggtt taaaagatca 720
tactcataca acgacaatgt agactttaag tgcaagtacg gatataaact atctggttcc 780
tcatcatcta cttgctctcc aggaaataca tggaagccgg aacttccaaa atgtgtacgc 840
<210> 3
<211> 1205
<212> DNA
<213> Artificial Sequence
<223> C-terminus (nucleotides 1786-3207) of amyloid
     precursor protein (APP)
<400> 3
taaaaaacgt ctaggccccc cgaaccacgg ggacgtggtt ttcctttgaa aaacacgata 60
ataccatgga tgcagaattc cgacatgact caggatatga agttcatcat caaaaattgg 120
tqttctttqc agaagatgtg ggttcaaaca aaggtgcaat cattggactc atggtgggcg 180
qtqttqtcat agcgacagtg atcgtcatca ccttggtgat gctgaagaag aaacagtaca 240
catccattca tcatggtgtg gtggaggttg acgccgctgt caccccagag gagcgccacc 300
tgtccaagat gcagcagaac ggctacgaaa atccaaccta caagttcttt gagcagatgc 360
aqaactagac ccccgccaca gcagcctctg aagttggaca gcaaaaccat tgcttcacta 420
cccatcggtg tccatttata gaataatgtg ggaagaaaca aacccgtttt atgatttact 480
cattatcgcc ttttgacagc tgtgctgtaa cacaagtaga tgcctgaact tgaattaatc 540
cacacatcag taatgtattc tatctctctt tacattttgg tctctatact acattattaa 600
tgggttttgt gtactgtaaa gaatttagct gtatcaaact agtgcatgaa tagattctct 660
cctgattatt tatcacatag ccccttagcc agttgtatat tattcttgtg gtttgtgacc 720
caattaagtc ctactttaca tatgctttaa gaatcgatgg gggatgcttc atgtgaacgt 780
ttaaacattt ttaagtattt cagatgcttt agagagattt tttttccatg actgcatttt 900
actgtacaga ttgctgcttc tgctatattt gtgatatagg aattaagagg atacacacgt 960
ttgtttcttc gtgcctgttt tatgtgcaca cattaggcat tgagacttca agcttttctt 1020
```

cttttacggg gcgggtgggg aggggtgctc tgctggtcga cgatccggct gctaacaaag 1140

cccgaaagga agctgagttg gctgctgcca ccgctgagca ataactagca taaccccttg 1200 gggcc <210> 4 <211> 100 <212> PRT <213> Artificial Sequence <223> C-terminus (nucleotides 1786-3207) of amyloid precursor protein (APP) Met Asp Ala Glu Phe Arg His Asp Ser Gly Tyr Glu Val His His Gln 10 Lys Leu Val Phe Phe Ala Glu Asp Val Gly Ser Asn Lys Gly Ala Ile Ile Gly Leu Met Val Gly Gly Val Val Ile Ala Thr Val Ile Val Ile 40 Thr Leu Val Met Leu Lys Lys Lys Gln Tyr Thr Ser Ile His His Gly 55 Val Val Glu Val Asp Ala Ala Val Thr Pro Glu Glu Arg His Leu Ser 70 75 Lys Met Gln Gln Asn Gly Tyr Glu Asn Pro Thr Tyr Lys Phe Phe Glu Gln Met Gln Asn 100 <210> 5 <211> 244 <212> PRT <213> Vaccinia Virus, Copenhagen Strain Cys Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Lys Phe Lys Asn Ser 10 Val Glu Thr Asp Ala Asn Ala Asn Tyr Asn Ile Gly Asp Thr Ile Glu 25 Tyr Leu Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr 40 Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys 55 Arg Arg Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly Gln Leu Asp Ile 70 75 Gly Gly Val Asp Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly 90 Tyr His Leu Ile Gly Glu Ser Lys Ser Tyr Cys Glu Leu Gly Ser Thr 105 Gly Ser Met Val Trp Asn Pro Glu Ala Pro Ile Cys Glu Ser Val Lys 120 Cys Gln Ser Pro Pro Ser Ile Ser Asn Gly Arg His Asn Gly Tyr Glu 135 140 Asp Phe Tyr Thr Asp Gly Ser Val Val Thr Tyr Ser Cys Asn Ser Gly 155 150 Tyr Ser Leu Ile Gly Asn Ser Gly Val Leu Cys Ser Gly Gly Glu Trp 170 165

Ser Asp Pro Pro Thr Cys Gln Ile Val Lys Cys Pro His Pro Thr Ile 180 185 190 Ser Asn Gly Tyr Leu Ser Ser Gly Phe Lys Arg Ser Tyr Ser Tyr Asn

200 195 Asp Asn Val Asp Phe Lys Cys Lys Tyr Gly Tyr Lys Leu Ser Gly Ser 215 Ser Ser Ser Thr Cys Ser Pro Gly Asn Thr Trp Lys Pro Glu Leu Pro 230 235 Lys Cys Val Arg <210> 6 <211> 244 <212> PRT <213> Vaccinia Virus, Western Reserve Strain Cys Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Lys Phe Lys Asn Ser Val Glu Thr Asp Ala Asn Ala Asn Tyr Asn Ile Gly Asp Thr Ile Glu Tyr Leu Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr 40 Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys Arg Arg Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly Gln Leu Asp Ile 70 75 Gly Gly Val Asp Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly 90 85 Tyr His Leu Ile Gly Glu Ser Lys Ser Tyr Cys Glu Leu Gly Ser Thr 105 Gly Ser Met Val Trp Asn Pro Glu Ala Pro Ile Cys Glu Ser Val Lys 120 Cys Gln Ser Pro Pro Ser Ile Ser Asn Gly Arg His Asn Gly Tyr Glu 135 Asp Phe Tyr Thr Asp Gly Ser Val Val Thr Tyr Ser Cys Asn Ser Gly 155 150 Tyr Ser Leu Ile Gly Asn Ser Gly Val Leu Cys Ser Gly Glu Trp 170 165 Ser Asp Pro Pro Thr Cys Gln Ile Val Lys Cys Pro His Pro Thr Ile 185 Ser Asn Gly Tyr Leu Ser Ser Gly Phe Lys Arg Ser Tyr Ser Tyr Asn 200 Asp Asn Val Asp Phe Lys Cys Lys Tyr Gly Tyr Lys Leu Ser Gly Ser 215 220 Ser Ser Ser Thr Cys Ser Pro Gly Asn Thr Trp Lys Pro Glu Leu Pro 235 230 Lys Cys Val Arg <210> 7 <211> 240 <212> PRT <213> Cowpox Virus, Russian Isolate From Human Patient Cys Cys Pro Ile Pro Ser Arg Pro Ile Thr Met Lys Phe Lys Gly Thr 10 Val Asp Ser His Tyr Asn Ile Gly Asp Thr Ile Glu Tyr Leu Cys Leu 25

Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys Arg Arg Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly Gln Leu Asp Ile Gly Gly Val Asp 70 Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly Tyr His Leu Ile Gly Glu Ser Lys Ser Tyr Cys Glu Leu Gly Ser Thr Gly Ser Met Val 105 Trp Asn Pro Glu Ala Pro Ile Cys Glu Ser Val Lys Cys Gln Ser Pro 120 Pro Ser Ile Ser Asn Gly Arg His Asn Gly Tyr Glu Asp Phe Tyr Thr 135 140 Asp Gly Ser Val Val Thr Tyr Ser Cys Asn Ser Gly Tyr Ser Leu Ile 150 155 Gly Asn Ser Gly Val Leu Cys Ser Gly Gly Glu Trp Ser Asp Pro Pro 170 Thr Cys Gln Ile Val Lys Cys Pro His Pro Thr Ile Ser Asn Gly Tyr 185 Leu Ser Ser Gly Phe Lys Arg Ser Tyr Ser Tyr Asn Asp Asn Val Asp 200 Phe Lys Cys Lys Tyr Gly Tyr Lys Leu Ser Gly Ser Ser Ser Thr 215 220 Cys Ser Pro Gly Asn Thr Trp Gln Pro Glu Leu Pro Lys Cys Val Arg 230

<211> 244 <212> PRT <213> Cowpox Virus, Brighton Strain <400> 8 Cys Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Lys Phe Lys Asn Ser 10 Val Gly Thr Asp Ala Asn Ala Asn Tyr Asn Ile Gly Asp Thr Ile Glu Tyr Leu Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys 55 Arg Lys Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly Gln Ile Asp Ile 70 75 Gly Gly Val Glu Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly Tyr Gln Leu Ile Gly Glu Ser Lys Ser Tyr Cys Glu Leu Gly Tyr Thr Gly Ser Met Val Trp Asn Pro Glu Ala Pro Ile Cys Glu Ser Val Lys 120 Cys Pro Ser Pro Pro Ser Val Thr Asn Gly Arg His Asn Gly Tyr Glu 135 Asp Phe Tyr Thr Asp Gly Ser Val Val Thr Tyr Ser Cys Asn Ser Gly 150 155 Tyr Ser Leu Ile Gly Asn Ser Gly Ile Val Cys Ser Gly Gly Glu Trp 170 165 Ser Asp Pro Pro Thr Cys Gln Ile Val Lys Cys Pro His Pro Ser Ile 185 180

<210> 8

Thr Asn Gly Tyr Leu Ser Ser Gly Phe Lys Arg Ser Tyr Ser His Asn 195 200 205

Asp Asn Val Asp Phe Lys Cys Arg His Gly Tyr Lys Leu Ser Gly Ser 210 215 220

Ser Ser Ser Thr Cys Ser Pro Gly Asn Thr Trp Gln Pro Glu Leu Pro 225 230 235 240

Lys Cys Val Arg

<210> 9 <211> 244 <212> PRT <213> Variola Virus, Bangladesh Strain Cys Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Lys Phe Lys Asn Ser Val Glu Thr Asp Ala Asn Ala Asn Tyr Asn Ile Gly Asp Thr Ile Glu 25 Tyr Leu Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr 40 Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys 55 Arg Arg Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly His Leu Asp Ile 70 Gly Gly Val Asp Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly 90 85 Tyr Tyr Leu Ile Gly Glu Tyr Lys Ser Tyr Cys Lys Leu Gly Ser Thr 105 Gly Ser Met Val Trp Asn Pro Lys Ala Pro Ile Cys Glu Ser Val Lys 120 Cys Gln Leu Pro Pro Ser Ile Ser Asn Gly Arg His Asn Gly Tyr Asn 135 Asp Phe Tyr Thr Asp Gly Ser Val Val Thr Tyr Ser Cys Asn Ser Gly 150 155 Tyr Ser Leu Ile Gly Asn Ser Gly Val Leu Cys Ser Gly Gly Glu Trp 170 Ser Asn Pro Pro Thr Cys Gln Ile Val Lys Cys Pro His Pro Thr Ile 180 185 Leu Asn Gly Tyr Leu Ser Ser Gly Phe Lys Arg Ser Tyr Ser Tyr Asn 200 Asp Asn Val Asp Phe Thr Cys Lys Tyr Gly Tyr Lys Leu Ser Gly Ser 215 Ser Ser Ser Thr Cys Ser Pro Gly Asn Thr Trp Gln Pro Glu Leu Pro 230 235 Lys Cys Val Arg

<210> 10
<211> 244
<212> PRT
<213> Variola Major Virus, Indian Strain
<400> 10
Cys Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Thr Phe Lys Asn Ser
1 5 10 15
Val Glu Thr Asp Ala Asn Ala Asn Tyr Asn Ile Gly Asp Thr Ile Glu

Tyr Leu Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys 55 Arg Arg Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly His Leu Asp Ile Gly Gly Val Asp Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly 90 Tyr Tyr Leu Ile Gly Glu Tyr Lys Ser Tyr Cys Lys Leu Gly Ser Thr 100 105 Gly Ser Met Val Trp Asn Pro Lys Ala Pro Ile Cys Glu Ser Val Lys 120 Cys Gln Leu Pro Pro Ser Ile Ser Asn Gly Arg His Asn Gly Tyr Asn 135 Asp Phe Tyr Thr Asp Gly Ser Val Val Thr Tyr Ser Cys Asn Ser Gly 150 Tyr Ser Leu Ile Gly Asn Ser Gly Val Leu Cys Ser Gly Gly Glu Trp 170 165 Ser Asn Pro Pro Thr Cys Gln Ile Val Lys Cys Pro His Pro Thr Ile 185 Leu Asn Gly Tyr Leu Ser Ser Gly Phe Lys Arg Ser Tyr Ser Tyr Asn 200 Asp Asn Val Asp Phe Thr Cys Lys Tyr Gly Tyr Lys Leu Ser Gly Ser 215 Ser Ser Ser Thr Cys Ser Pro Gly Asn Thr Trp Gln Pro Glu Leu Pro 235 230 Lys Cys Val Arg

```
<210> 11
<211> 244
<212> PRT
<213> Variola Minor Virus, Alastrim Garcia Strain
Cys Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Lys Phe Lys Asn Ser
                                    10
Val Glu Thr Asp Ala Asn Ala Asn Tyr Asn Ile Gly Asp Thr Ile Glu
                                25
            20
Tyr Leu Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr
                            40
Ala Lys Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys
                        55
Arg Arg Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly His Leu Asp Ile
                                        75
Gly Gly Val Asp Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly
                                    90
Tyr Tyr Leu Ile Gly Glu Tyr Lys Ser Tyr Cys Lys Leu Gly Ser Thr
                                105
Gly Ser Met Val Trp Asn Pro Lys Ala Pro Ile Cys Glu Ser Val Lys
                            120
Cys Gln Leu Pro Pro Ser Ile Ser Asn Gly Arg His Asn Gly Tyr Asn
                        135
                                            140
Asp Phe Tyr Thr Asp Gly Ser Val Val Thr Tyr Ser Cys Asn Ser Gly
                                        155
                    150
Tyr Ser Leu Ile Gly Asn Ser Gly Val Leu Cys Ser Gly Gly Glu Trp
```

 Ser Asn Pro Pro Pro Thr Cys Gln Ile Val Lys Cys Pro Tyr Pro Thr Ile 180
 185
 185
 190
 190
 186
 190
 185
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190
 190

<210> 12 <211> 197 <212> PRT <213> Monkeypox Virus, Isolated from a Human Patient fr6 <400> 12 Tyr Cys Thr Ile Pro Ser Arg Pro Ile Asn Met Lys Phe Lys Asn Ser Val Glu Thr Asp Ala Asn Tyr Asn Ile Gly Asp Thr Ile Glu Tyr Leu 25 Cys Leu Pro Gly Tyr Arg Lys Gln Lys Met Gly Pro Ile Tyr Ala Lys 40 Cys Thr Gly Thr Gly Trp Thr Leu Phe Asn Gln Cys Ile Lys Arg Arg 55 Cys Pro Ser Pro Arg Asp Ile Asp Asn Gly Gln Leu Asp Ile Gly Gly 70 Val Asp Phe Gly Ser Ser Ile Thr Tyr Ser Cys Asn Ser Gly Tyr His 90 Leu Ile Gly Glu Ser Lys Ser Tyr Cys Glu Leu Gly Ser Thr Gly Ser 100 105 Met Val Trp Asn Pro Glu Ala Pro Ile Cys Glu Ser Val Lys Cys Gln 120 Ser Pro Pro Ser Ile Ser Asn Gly Arg His Asn Gly Tyr Glu Asp Phe 135 140 Tyr Ile Asp Gly Ser Ile Val Thr Tyr Ser Cys Asn Ser Gly Tyr Ser 150 155 Leu Ile Gly Asn Ser Gly Val Met Cys Ser Gly Glu Trp Ser Asn 165 170 Pro Pro Thr Cys Gln Ile Val Lys Cys Pro His Pro Ile Ser Asn Gly 185 180 Lys Leu Leu Ala Ala

195